Considerd DT 5/2/05

PATENT

ANM243.1APCI

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

pplicant

Gabant et al.

Appl. No.

: 10/031,021

Filed

: March 19, 2002

For

: NON-HUMA' GENETICALLY

MODIFIED MAMMAL LACKING

THE ALPHA PETOPROTEIN

Examinet

Nguyen, D. T.

Group Art Unit

1632

DECLARA: JON UNDER 37 C.F.R. §1.132

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

## Dear Sir:

- 1. This Declaration is being submitted to demonstrate that the knowledge of how to create a modification in a mouse AFP gene using ES cell technology was readily available to a person skilled in the art at the time the Ui: Provisional Application No.: 60/143,269 to which this Application claims priority was filed.
- I am an inventor on the above-identified patent application and am familiar with the specification and prosecution history.
- 3. I have extensive experience in the field of the claimed invention as indicated in the attached Curriculum Vitae provided herewith as Exhibit A.
- 4. In some embodiments of the presently claimed mice, at least one allele of the endogenous genetic sequence of the wild-type alpha-fetoprotein (AFP) is disrupted to prevent the mice from making sufficient active AFP to confix fertility. When such a genetically modified mouse has just one allele of the AFP gene modified, it can be mated with another such mouse to obtain